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NEWS X25

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COST IN U.S. DOLLARS

ENTRY SESSION FULL ESTIMATED COST 0.21 0.21

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FILE 'EMBASE' ENTERED AT 11:02:22 ON 21 FEB 2007

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FILE 'BIOSIS' ENTERED AT 11:02:22 ON 21 FEB 2007

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=> s vgf or vgfarp

L1838 VGF OR VGFARP

=> s l1 and (alzheimer or alsheimer)

30 L1 AND (ALZHEIMER OR ALSHEIMER)

=> dup rem 12

PROCESSING COMPLETED FOR L2

20 DUP REM L2 (10 DUPLICATES REMOVED)

=> dis ibib abs 13 1-20

ANSWER 1 OF 20 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2007:63555 CAPLUS

DOCUMENT NUMBER:

146:138265

TITLE:

SELDI mass spectrometry and immunoassay methods for

SINCE FILE

TOTAL

determination of VGF peptide-1 as a biomarker for

Alzheimer's disease

INVENTOR (S):

Davies, Huw Alun; Blennow, Kaj; McGuire, James;

Podust, Vladimir; Simonsen, Anja Hviid

PATENT ASSIGNEE(S):

Ciphergen Biosystems, Inc., USA

SOURCE:

U.S. Pat. Appl. Publ., 19pp., Cont.-in-part of U.S.

Ser. No. 982,545.

CODEN: USXXCO

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
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US 2007015221	A1	20070118	US 2006-452477	20060613
US 2005244890	Al	20051103	US 2004-982545	20041106
WO 2006113289	A2	20061026	WO 2006-US13727	20060411
W: AE, AG, AL,	AM, AT	, AU, AZ, BA	A, BB, BG, BR, BW, BY,	BZ, CA, CH,
. CN, CO, CR,	CU, CZ	, DE, DK, DM	1, DZ, EC, EE, EG, ES,	FI, GB, GD,

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PRIORITY APPLN. INFO.:
                                                 US 2004-572617P
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                                                                           20040518
                                                 US 2004-586503P
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                                                                           20040708
                                                 US 2004-982545
                                                                        A2 20041106
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                                                                           20050419
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                                                                           20050616
                                                 WO 2006-US13727
                                                                        A2 20060411
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                                                 US 2003-526753P
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                                                                           20040219
                                                 US 2004-547250P
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                                                                           20040223
                                                 US 2004-558896P
                                                                        Р
                                                                           20040402
     The present invention provides a neurosecretory protein VGF
AB
     peptide useful in qualifying Alzheimer's disease status in a
     patient. In particular, this peptide and modified forms thereof may be
     used to classify a subject sample as Alzheimer's disease or non-
     Alzheimer's disease. The peptide biomarker can be detected by
     SELDI mass spectrometry.
     ANSWER 2 OF 20 CAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER:
                            2006:1357056 CAPLUS
DOCUMENT NUMBER:
                            146:58251
TITLE:
                            Fragment of neurosecretory protein VGF as a
                            biomarker for Alzheimer's disease
INVENTOR(S):
                            Davies, Huw Alun; Blennow, Kaj; McGuire, James;
                            Podust, Vladimir; Simonsen, Anja Hviid
PATENT ASSIGNEE(S):
                            Ciphergen Biosystems, Inc., USA
                            PCT Int. Appl., 43pp.
SOURCE:
                            CODEN: PIXXD2
DOCUMENT TYPE:
                            Patent
LANGUAGE:
                            English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
                            KIND
                                    DATE
                                                 APPLICATION NO.
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     WO 2006138325
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                                               WO 2006-US23044
                                                                           20060613
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              GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
              KG, KZ, MD, RU, TJ, TM
PRIORITY APPLN. INFO.:
                                                 US 2005-691637P
                                                                        P 20050616
AB
     The present invention provides SELDI mass spectrometry and immunoassay
     methods for the determination of a neurosecretory protein VGF peptide-1,
     which useful in qualifying Alzheimer's disease status in a
```

patient. In particular, this peptide and modified forms thereof may be

used to classify a subject sample as Alzheimer's disease or non-

Alzheimer's disease.

L3 ANSWER 3 OF 20 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2006:1062718 CAPLUS

DOCUMENT NUMBER: 145:416031

TITLE: Biological fluid markers for diagnosis and monitoring

of neurodegenerative disease

INVENTOR(S): Schulman, Howard; Lowe, David; Becker, Christopher H.;

Zhou, Haihong; Roy, Sushmita Mimi

PATENT ASSIGNEE(S): Neurodx, LLC, USA SOURCE: PCT Int. Appl., 161pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PAT	PATENT NO.				KIND		DATE		APPLICATION NO.						DATE		
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WO	2006	1080	51		A2		2006	1012	Ī	WO 2	006-1	JS12	581		2	0604	405
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		IS,	IT,	LT,	LU,	LV,	MC,	NL,	PL,	PT,	RO,	SE,	SI,	SK,	TR,	BF,	ВJ,
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		KG,	ΚZ,	MD,	RU,	ΤJ,	TM										
ORITY	APP	LN.	INFO	.:					1	US 2	005-	66824	45P	1	P 2	0050	405

PRIO The present invention provides compns., methods and kits useful for the AB diagnosis and treatment of neurodegenerative diseases, e.g., Alzheimer's disease (AD). In particular, the invention provides polypeptides and metabolites that are markers of AD, polynucleotides that encode the polypeptides, and antibodies that specifically bind to the polypeptides. The invention also provides methods for using the polypeptides, metabolites, polynucleotides and antibodies in the diagnosis and treatment of AD, monitoring progression of the disease and screening of candidate therapeutic compds. Thus, certain proteins and metabolites were found to be differentially expressed in cerebrospinal fluid (CSF) samples from AD subjects and subjects with mild cognitive impairment compared to CSF samples from control subjects. The CSF samples were separated into high mol. wt (>5 kDa) and low mol. weight fractions. After removal of high abundance proteins, the high mol. weight sample was digested with trypsin, separated by chromatog., and analyzed by mass spectrometry.

L3 ANSWER 4 OF 20 EMBASE COPYRIGHT (c) 2007 Elsevier B.V. All rights reserved on STN DUPLICATE 1

ACCESSION NUMBER: 2006599527 EMBASE

TITLE: Disease biomarkers in cerebrospinal fluid of patients with

first-onset psychosis.

AUTHOR: Huang J.T.-J.; Leweke F.M.; Oxley D.; Wang L.; Harris N.;

Koethe D.; Gerth C.W.; Nolden B.M.; Gross S.; Schreiber D.;

Reed B.; Bahn S.

CORPORATE SOURCE: S. Bahn, Institute of Biotechnology, University of

Cambridge, Cambridge, United Kingdom. sb209@cam.ac.uk

SOURCE: PLoS Medicine, (2006) Vol. 3, No. 11, pp. 2145-2158. .

Refs: 32

ISSN: 1549-1277 E-ISSN: 1549-1676

COUNTRY: United States
DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 029 Clinical Biochemistry

032 Psychiatry

LANGUAGE: English
SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 12 Jan 2007

Last Updated on STN: 12 Jan 2007

Background: Psychosis is a severe mental condition that is characterized AB by a loss of contact with reality and is typically associated with hallucinations and delusional beliefs. There are numerous psychiatric conditions that present with psychotic symptoms, most importantly schizophrenia, bipolar affective disorder, and some forms of severe depression referred to as psychotic depression. The pathological mechanisms resulting in psychotic symptoms are not understood, nor is it understood whether the various psychotic illnesses are the result of similar biochemical disturbances. The identification of biological markers (so-called biomarkers) of psychosis is a fundamental step towards a better understanding of the pathogenesis of psychosis and holds the potential for more objective testing methods. Methods and Findings: Surface-enhanced laser desorption ionization mass spectrometry was employed to profile proteins and peptides in a total of 179 cerebrospinal fluid samples (58 schizophrenia patients, 16 patients with depression, five patients with obsessive-compulsive disorder, ten patients with Alzheimer disease, and 90 controls). Our results show a highly significant differential distribution of samples from healthy volunteers away from drug-naive patients with first-onset paranoid schizophrenia. The key alterations were the up-regulation of a 40-amino acid VGF -derived peptide, the down-regulation of transthyretin at .apprx.4 kDa, and a peptide cluster at .apprx.6,800-7,300 Da (which is likely to be influenced by the doubly charged ions of the transthyretin protein cluster). These schizophrenia-specific protein/peptide changes were replicated in an independent sample set. Both experiments achieved a specificity of 95% and a sensitivity of 80% or 88% in the initial study and in a subsequent validation study, respectively. Conclusions: Our results suggest that the application of modern proteomics techniques, particularly mass spectrometric approaches, holds the potential to advance the understanding of the biochemical basis of psychiatric disorders and may in turn allow for the development of diagnostics and improved therapeutics. Further studies are required to validate the clinical effectiveness and disease specificity of the identified biomarkers. Copyright: .COPYRGT. 2006 Huang et al.

L3 ANSWER 5 OF 20 MEDLINE on STN ACCESSION NUMBER: 2006695642 MEDLINE DOCUMENT NUMBER: PubMed ID: 17090210

TITLE: Disease biomarkers in cerebrospinal fluid of patients with

first-onset psychosis.

AUTHOR: Huang Jeffrey T-J; Leweke F Markus; Oxley David; Wang Lan;

Harris Nathan; Koethe Dagmar; Gerth Christoph W; Nolden Brit M; Gross Sonja; Schreiber Daniela; Reed Benjamin; Bahn

Sabine

CORPORATE SOURCE: Institute of Biotechnology, University of Cambridge,

Cambridge, United Kingdom.

SOURCE: PLoS medicine, (2006 Nov) Vol. 3, No. 11, pp. e428.

Journal code: 101231360. E-ISSN: 1549-1676.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

(RESEARCH SUPPORT, NON-U.S. GOV'T)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200701

ENTRY DATE: Entered STN: 30 Nov 2006

Last Updated on STN: 9 Jan 2007 Entered Medline: 8 Jan 2007

AB BACKGROUND: Psychosis is a severe mental condition that is characterized

hallucinations and delusional beliefs. There are numerous psychiatric conditions that present with psychotic symptoms, most importantly schizophrenia, bipolar affective disorder, and some forms of severe depression referred to as psychotic depression. The pathological mechanisms resulting in psychotic symptoms are not understood, nor is it understood whether the various psychotic illnesses are the result of similar biochemical disturbances. The identification of biological markers (so-called biomarkers) of psychosis is a fundamental step towards a better understanding of the pathogenesis of psychosis and holds the potential for more objective testing methods. METHODS AND FINDINGS: Surface-enhanced laser desorption ionization mass spectrometry was employed to profile proteins and peptides in a total of 179 cerebrospinal fluid samples (58 schizophrenia patients, 16 patients with depression, five patients with obsessive-compulsive disorder, ten patients with Alzheimer disease, and 90 controls). Our results show a highly significant differential distribution of samples from healthy volunteers away from drug-naive patients with first-onset paranoid schizophrenia. The key alterations were the up-regulation of a 40-amino acid VGF -derived peptide, the down-regulation of transthyretin at approximately 4 kDa, and a peptide cluster at approximately 6,800-7,300 Da (which is likely to be influenced by the doubly charged ions of the transthyretin protein cluster). These schizophrenia-specific protein/peptide changes were replicated in an independent sample set. Both experiments achieved a specificity of 95% and a sensitivity of 80% or 88% in the initial study and in a subsequent validation study, respectively. CONCLUSIONS: Our results suggest that the application of modern proteomics techniques, particularly mass spectrometric approaches, holds the potential to advance the understanding of the biochemical basis of psychiatric disorders and may in turn allow for the development of diagnostics and improved therapeutics. Further studies are required to validate the clinical effectiveness and disease specificity of the identified biomarkers.

by a loss of contact with reality and is typically associated with

ANSWER 6 OF 20 CAPLUS COPYRIGHT 2007 ACS on STN L3

ACCESSION NUMBER: 2005:451526 CAPLUS

DOCUMENT NUMBER: 143:5662

TITLE: Identification of biomarkers for Alzheimer's

disease by expression profiling and SELDI mass

spectrometry

Davies, Huw Alun; McGuire, James; Simonsen, Anja INVENTOR (S):

Hviid; Blennow, Kaj; Podust, Vladimir

Ciphergen Biosystems, Inc., USA PATENT ASSIGNEE(S):

SOURCE:

PCT Int. Appl., 145 pp. CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

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T-10	2005	0474					20050506			WO 2004-US37994					2.	2041	106	
WO	2005	04/4	84		AZ		2005	0526	1	WO Z	004-	US3/	<b>774</b>		21	JU41.	100	
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EP	1694	816			A2		2006	0830		EP 2	004 -	8109	48		2	0041	106	

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                HR, IS, YU
PRIORITY APPLN. INFO.:
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AB
      The present invention provides protein-based biomarkers and biomarker
      combinations that are useful in qualifying Alzheimer's disease
      status in a patient. In particular, the biomarkers of this invention are
      useful to classify a subject sample as Alzheimer's or non-
      Alzheimer's dementia or normal. The biomarkers can be detected by
      SELDI mass spectrometry. In addition, the invention provides appropriate
      treatment interventions and methods for measuring response to treatment.
      Certain biomarkers of the invention may also be suitable for employment as
      radio-labeled ligands in non-invasive imaging techniques such as Positron
      Emission Tomog. (PET).
      ANSWER 7 OF 20 CAPLUS COPYRIGHT 2007 ACS on STN
T.3
ACCESSION NUMBER:
                               2005:239024 CAPLUS
DOCUMENT NUMBER:
                               142:311677
                               Protein complexes associated with \beta-amyloid
TITLE:
                               precursor protein processing and their use for
                               diagnosis and therapy of Alzheimer's disease
                                and other neurodegeneration disorders
                                Bouwmeester, Tewis; Drewes, Gerard; Hopf, Carsten;
INVENTOR(S):
                                Joberty, Gerard; Rowley, Adele
PATENT ASSIGNEE(S):
                               Cellzome A.-G., Germany
                                PCT Int. Appl., 1294 pp.
SOURCE:
                                CODEN: PIXXD2
DOCUMENT TYPE:
                                Patent
LANGUAGE:
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FAMILY ACC. NUM. COUNT:
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RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
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                SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
                SN, TD, TG
      EP 1670903
                                        20060621
                                A2
                                                     EP 2004-764730
                                                                                   20040902
           R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
                IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK
PRIORITY APPLN. INFO.:
                                                      EP 2003-19642
                                                                               A 20030905
                                                      WO 2003-EP13980
                                                                               W 20031210
                                                      EP 2004-1894
                                                                               A 20040129
                                                      EP 2004-1895
                                                                              A 20040129
                                                      EP 2004-7447
                                                                              A 20040326
                                                      WO 2004-EP4889
                                                                              A 20040507
                                                      WO 2004-EP4891
                                                                              A 20040507
                                                      EP 2004-18874
                                                                               A 20040809
                                                                               W 20040902
                                                      WO 2004-EP9771
      The present invention relates to protein complexes of the \beta-amyloid
AB
      precursor protein (APP) processing pathway, component proteins of the said
      complexes, fragments and derivs. of the component proteins, and antibodies
      specific to the complexes. Thus, two-hybrid screening identified 266
      protein components among 14 protein complexes: the presentlin 1 complex,
      presenilin 2 complex, nicastrain complex, Aph-la complex, Aph-lb complex,
      Pen-2 complex, BACE (β-secretase) N215D complex, APP complex,
      APP695SW complex, APP-C99 complex, Tau complex, X11\beta complex, Fe65
      complex. and calsenilin complex. The present invention also relates to
      methods for use of the complexes of the APP processing pathway and their
      interacting proteins in, inter alia, screening, diagnosis, and therapy, as
      well as to methods of preparing the complexes.
REFERENCE COUNT:
                                      THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS
                               11
                                      RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
      ANSWER 8 OF 20 CAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER:
                               2005:158835 CAPLUS
DOCUMENT NUMBER:
                               142:234397
TITLE:
                               Microarray systems and methods for diagnosing and
                               treating psychol. and behavioral conditions, and
                               assessing efficacy of therapy, based on gene
                               expression signatures
INVENTOR(S):
                              Duman, Ronald; Sathyanesan, Samuel
                              Yale University, USA
PATENT ASSIGNEE(S):
                               PCT Int. Appl., 94 pp.
SOURCE:
                               CODEN: PIXXD2
DOCUMENT TYPE:
                               Patent
LANGUAGE:
                               English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:
      PATENT NO.
                                                      APPLICATION NO.
                              KIND
                                       DATE
                                                                                   DATE
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                              A2
      WO 2005017203
                                       20050224
                                                      WO 2004-US22178
                                                                                  20040712
                                     20050909
      WO 2005017203
                               А3
          2005017203

A3 20050909

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                SN, TD, TG
      US 2005084880
                                       20050421
                                                      US 2004-889336
                                A1
                                                                                   20040712
                                                      US 2003-486632P P 20030711
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The systems and methods described herein include microarray systems and

PRIORITY APPLN. INFO.:

methods for manufacturing and printing microarrays to provide gene chips capable

of detecting gene signatures of psychiatric conditions, and as well as gene chips and arrays of sequences for such applications. The invention further provides methods of identifying gene signatures for psychiatric conditions, methods of treating such conditions, and methods of identifying therapeutics for the treatment of neurol. and psychiatric conditions. The examples present gene expression profile data from brain tissue following electroconvulsive seizure (ECS) therapy, and define an ECS gene signature.

L3 ANSWER 9 OF 20 MEDLINE on STN DUPLICATE 2

ACCESSION NUMBER: 2005119860 MEDLINE DOCUMENT NUMBER: PubMed ID: 15749342

TITLE: Amyloid-beta-stimulated plasminogen activation by

tissue-type plasminogen activator results in processing of

neuroendocrine factors.

AUTHOR: Kranenburg O; Gent Y Y J; Romijn E P; Voest E E; Heck A J

R; Gebbink M F B G

CORPORATE SOURCE: Department of Medical Oncology, University Medical Center

Utrecht, Utrecht, The Netherlands.

SOURCE: Neuroscience, (2005) Vol. 131, No. 4, pp. 877-86.

Journal code: 7605074. ISSN: 0306-4522.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200505

ENTRY DATE: Entered STN: 8 Mar 2005

Last Updated on STN: 13 May 2005 Entered Medline: 12 May 2005

Alzheimer's disease brain is characterized by the abundant AB presence of amyloid deposits. Accumulation of the major constituent of these deposits, amyloid-beta (Abeta), has been associated with decreased neurotransmission, increased neuronal cell death, and with cognitive decline. The mechanisms underlying these phenomena have not yet been fully elucidated. We have previously shown that amyloid peptides like Abeta bind tissue-type plasminogen activator (tPA) and cause enhanced plasmin production. Here we describe the identification of five major neuronal cell-produced Abeta-associated proteins and how Abeta-stimulated plasmin formation affects their processing. These five proteins are all neuroendocrine factors (NEFs): chromogranins A, B and C; truncated chromogranin B; and VGF. Plasminogen caused processing of Abeta-bound (but not soluble) tPA, chromogranin B and VGF and the degradation products were released from Abeta. Processing of the neuroendocrine factors was dependent on tPA as it was largely abrogated in tPA-/- cells or in the presence of a specific tPA-inhibitor. If plasmin indeed produces NEF-derived peptides in vivo, some of these peptides may have biological activity, for instance in regulating neurotransmitter release that may affect the pathology of Alzheimer's disease.

L3 ANSWER 10 OF 20 MEDLINE on STN DUPLICATE 3

ACCESSION NUMBER: 2006078566 MEDLINE DOCUMENT NUMBER: PubMed ID: 16464167

TITLE: Identification of novel biomarker candidates by

differential peptidomics analysis of cerebrospinal fluid in

Alzheimer's disease.

AUTHOR: Selle Hartmut; Lamerz Jens; Buerger Katharina; Dessauer

Andreas; Hager Klaus; Hampel Harald; Karl Johann; Kellmann Markus; Lannfelt Lars; Louhija Jukka; Riepe Matthias; Rollinger Wolfgang; Tumani Hayrettin; Schrader Michael;

Zucht Hans-Dieter

CORPORATE SOURCE: BioVisioN AG, 30625 Hannover, Germany...

h.selle@peptidomics.de

Combinatorial chemistry & high throughput screening, (2005 SOURCE:

Dec) Vol. 8, No. 8, pp. 801-6.

Journal code: 9810948. ISSN: 1386-2073.

PUB. COUNTRY:

Netherlands

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

200603

ENTRY DATE:

Entered STN: 9 Feb 2006

Last Updated on STN: 8 Mar 2006 Entered Medline: 7 Mar 2006

The objective of this work was the application of peptidomics technologies AΒ for the detection and identification of reliable and robust biomarkers for Alzheimer's disease (AD) contributing to facilitate and further improve the diagnosis of AD. Using a new method for the comprehensive and comparative profiling of peptides, the differential peptide display (DPD), 312 cerebrospinal fluid (CSF) samples from AD patients, cognitively unimpaired subjects and from patients suffering from other primary dementia disorders were analysed as four independent analytical sets. By combination with a cross validation procedure, candidates were selected from a total of more than 6,000 different peptide signals based on their discriminating power. Twelve candidates were identified using mass-spectrometric techniques as fragments of the possibly neuroprotective neuroendocrine protein VGF and another one as the complement factor C3 descendent C3f. The combination of peptide profiling and cross validation resulted in the detection of novel potential biomarkers with

ANSWER 11 OF 20 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on L3

remarkable robustness and a close relation to AD pathophysiology.

STN

ACCESSION NUMBER: DOCUMENT NUMBER:

2005:372737 BIOSIS PREV200510171680

TITLE:

Amyloid-beta-stimulated plasminogen activation by tPA

results in processing of neuroendocrine factors.

AUTHOR(S):

Kranenburg, O. [Reprint Author]; Gent, Y. Y. J.; Romijn, E.

P:; Voest, E. E.; Heck, A. J. K.; Gebbink, M. F. B. G.

CORPORATE SOURCE: SOURCE:

Univ Utrecht, Med Ctr, Dept Med Oncol, Utrecht, Netherlands Thrombosis and Haemostasis, (APR 2005) Vol. 93, No. 4, pp.

Meeting Info.: 10th Interenational Workshop on Molecular and Cellular Biology of Plasminogen Activation. Washington,

DC, USA. April 09 -13, 2005. CODEN: THHADQ. ISSN: 0340-6245.

DOCUMENT TYPE:

Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

LANGUAGE:

English

ENTRY DATE:

Entered STN: 21 Sep 2005

Last Updated on STN: 21 Sep 2005

ANSWER 12 OF 20 CAPLUS COPYRIGHT 2007 ACS on STN  $L_3$ 

ACCESSION NUMBER: DOCUMENT NUMBER:

2004:1081026 CAPLUS

142:50129

TITLE:

Microarray for determining expression of

psychoneuroendocrinimmune genes and diagnosis of

diseases

INVENTOR(S):

Nicholson, Ainsley; Vernon, Suzanne D.

PATENT ASSIGNEE(S):

The Government of the United States as Represented by the Secretary of the Department of Health and Human

SOURCE:

Services, USA PCT Int. Appl., 254 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

## PATENT INFORMATION:

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DATE
                                                                                                         APPLICATION NO.
           PATENT NO.
                                                           KIND
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                                                                                                    WO 2004-US17686
           WO 2004108899
                   2004108899

A2 20041216 WO 2004-US17686 20040604
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
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LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
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SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
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                                                                           20041216
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                               SN, TD, TG
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           CA 2528162
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PRIORITY APPLN. INFO.:
                                                                                                          US 2003-475915P
                                                                                                                                                   W 20040604
                                                                                                           WO 2004-US17686
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Disclosed are compns. and methods for microarrays comprising genes AB involved in psychoneuroendocrinimmune (PNI) activity. An oligonucleotide microarray composed entirely of PNI genes was designed, which can allow a researcher to assess the overall psychoneuroendocrineimmune state of an individual, and to observe systemic responses to various stresses. The PNI array has widespread applicability and marketability in the diagnosis and treatment of diseases that result from dysregulation of the hypothalamic-pituitary-adrenal axis. A total of 1451 genes encoding 1738 transcriptional products can be distinguished and samples from human or mouse can hybridize with equal affinity, facilitating animal studies. Arabidopsis and housekeeping genes are used as controls. To determine the extent of peripheral blood PNI gene expression, both EST and microarray databases were queried; there were 566 genes from an EST database that matched to one of 1622 genes in the PNI database. The utility of the PNI array is demonstrated for research of chronic fatigue syndrome and other diseases involving PNI.

ANSWER 13 OF 20 CAPLUS COPYRIGHT 2007 ACS on STN

2004:183119 CAPLUS ACCESSION NUMBER:

DOCUMENT NUMBER:

140:232100

TITLE:

SELDI-TOF MS detection and identification of protein

biomarkers for diagnosing Alzheimer's

INVENTOR(S):

Yalkinoglu, Oezkan; Koenig, Gerhard; Hochstrasser,

Denis Francois; Sanchez, Jean-Charles; Carrette, Odile

PATENT ASSIGNEE(S): Bayer Healthcare A.-G., Germany

SOURCE:

PCT Int. Appl., 38 pp.

DOCUMENT TYPE:

CODEN: PIXXD2 Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 2

PATENT	NO.			KIN	D	DATE APPLICATION NO.						D	DATE			
<b>-</b>					_											
WO 2004	01904	43		A2		2004	0304	1	WO 2	003-1	EP88'	79		20	0030	811
WO 2004	01904	43		<b>A3</b>		2004	0624									
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	TR,	TT,	TZ,	UΑ,	ÜĠ,	US,	UZ,	VC,	VN,	YU,	ZA,	ZM,	zw			
RW:	GH.	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	ΰĠ,	ZM,	ZW,	AM,	AZ,	ΒY,

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      EP 1394549
                                 A1
                                         20040303
                                                        EP 2002-18283
                                                                                      20020823
           R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK
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                                                                                      20030811
      EP 1535076
                                 A2
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                                                        EP 2003-792291
                                                                                      20030811
           R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
      JP 2005536729
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                                         20051202
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                                                                                      20030811
      US 2006240561
                                 A1
                                         20061026
                                                        US 2006-525633
                                                                                      20060410
PRIORITY APPLN. INFO.:
                                                        EP 2002-18283
                                                                                  A 20020823
                                                        EP 2002-26643
                                                                                  A 20021129
                                                        WO 2003-EP8879
                                                                                  W
                                                                                      20030811
      A method for assessing the state of Alzheimer's disease in
AB
      patients is disclosed. A method for monitoring the progression of
      Alzheimer's disease in patients is also disclosed. The method applies detection of specific peptide markers, e.g., using mass
      spectrometric anal. (SELDI-TOF MS). The specific markers are: human cystatin C, human \beta-2-microglobulin, human myoglobin (new variant),
      neurosecretory protein VGF or fragments of these proteins.
      Protein chip SELDI anal. of CSF on SAX2 chip is described. In order to
      identify the protein markers observed by SELDI, a fractionation of crude CSF
      on strong anionic exchange chromatog. column was performed. The proteins
      were further purified by gel electrophoresis and identified by peptide
      mass fingerprinting anal. and peptide fragmentation anal.
      ANSWER 14 OF 20 CAPLUS COPYRIGHT 2007 ACS on STN
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ACCESSION NUMBER: 2004:60633 CAPLUS

DOCUMENT NUMBER: 140:126705

TITLE: Markers of neuronal cell death and their use in

diagnosis and therapy

INVENTOR(S): Zack, Donald J.; Kageyama, Masaaki PATENT ASSIGNEE(S): The Johns Hopkins University, USA

PCT Int. Appl., 109 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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PATENT NO.
                        KIND
                                DATE
                                            APPLICATION NO.
                                                                   DATE
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    WO 2004007673
                         A2
                                20040122
                                            WO 2003-US21729
                                                                   20030714
    WO 2004007673
                         A3
                                20041118
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
             PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN,
             TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
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    AU 2003249054
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                                20040202
                                           AU 2003-249054
                                                                   20030714
    US 2004086511
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                                20040506
                                            US 2003-617885
                                                                   20030714
PRIORITY APPLN. INFO.:
                                                                P 20020712
                                            US 2002-395753P
                                            WO 2003-US21729
                                                                W 20030714
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Neuronal cell death, as modeled by removal of serum or NGF from growth AB medium, is characterized by many changes in gene expression. Gene expression was compared before and after withdrawal of serum or NGF.

These results provide clues to underlying mol. processes occurring during neuronal and photoreceptor degeneration, and provide direction for future cell-based studies.

L3 ANSWER 15 OF 20 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2004:2706 CAPLUS

DOCUMENT NUMBER: 140:53449

TITLE: Pharmaceutical compositions for the treatment of

diseases related to neurotrophins

INVENTOR(S): Guarna, Antonio; Cozzolino, Federico; Torcia, Maria;

Garaci, Enrico

PATENT ASSIGNEE(S): Italy

SOURCE: PCT Int. Appl., 76 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE: Eng FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.						KIND DATE				APP	LICAT	ION I	NO.	DATE			
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												, ZA,		•		,		•
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			FI,	FR,	GB,	GR,	ΗU,	IE,	IT,	LU,	MC	, NL,	PT,	RO,	SE,	SI,	SK,	TR,
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	IT	2002	FI01	07		A1		2003	1219		IT	2002-	FI10	7		2	0020	619
	CA	2489	965			A1		2003	1231		CA	2003-	2489	965		2	0030	618
	AU	2003	2465	59		Al		2004	0106		AU	2003-	2465	59		2	0030	618
	EΡ	1551	412			, Al		2005	0713		EΡ	2003-	7606	52		2	0030	618
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		1662										2003-						
												2004-					0030	618
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PRIC	RITY	Y APP	LN.	INFO	. :						IT	2002-	FI10	7		A 2	0020	619
										_	WO	2003-	EP64	71	1	W 2	0030	618
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OTHER SOURCE(S): MARPAT 140:53449

The invention refers to pharmaceutical prepns. including as active compds. 3-aza-bicyclo[3.2.1]octane derivs. and/or their dimers acting as agonists of human neurotrophins. Therefore, such compds. are useful for treatment of diseases in which the neurotrophin functions are involved in defect, particularly of Nerve Growth Factor (NGF), such as neurodegenerative diseases of central nervous system (CNS), acquired immunodeficiency due to a reduced NGF bioavailability, or morbous conditions in which the stimulus of neoangiogenesis process is convenient.

REFERENCE COUNT:

THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 16 OF 20 CAPLUS COPYRIGHT 2007 ACS on STN

3

ACCESSION NUMBER: 2003:242437 CAPLUS

DOCUMENT NUMBER: 138:249938

---- 136:249936

TITLE: Gene expression profile biomarkers and therapeutic targets for brain aging and age-related cognitive

impairment in rats

INVENTOR(S): Landfield, Philip W.; Blalock, Eric M.; Chen,

Kuey-Chu; Foster, Thomas C.

PATENT ASSIGNEE(S): University of Kentucky Research Foundation, USA

SOURCE: PCT Int. Appl., 84 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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                                       20030327 WO 2002-US25607
     WO 2003025122
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PRIORITY APPLN. INFO.:
                                                      US 2001-311343P
                                                                                P 20010813
     A statistical and functional correlation strategy is provided to identify
      changes in cellular pathways specifically linked to impaired cognitive
      function with aging. Analyses using the strategy identified multiple
     groups of genes expressed in the hippocampal CA1 region of rats, where the
     genes were expressed at different levels for several ages. The aging
     changes in expression began before mid-life. Many of the genes were
      involved in specific neuronal and glial pathways with previously
     unrecognized relationships to aging and/or cognitive decline. The
     processes identified by the strategy suggest a new hypothesis of brain
     aging in which initially decreased neuronal activity and/or oxidative
     metabolism trigger sep. but parallel genomic cascades in neurons and glia.
     neurons, the cascade results in elevations in calcium signaling and redns.
     of immediate early gene signaling, biosynthesis, synaptogenesis, and
     neurite remodeling. In contrast, glia undergo increased lipid metabolism and
     mediate a cycle of demyelination and remyelination that induces antigen
     presentation, inflammation, oxidative stress, and extracellular
     restructuring. These identified genes and the proteins they encode can be
     used as novel biomarkers of brain aging and as targets for developing
      treatment methods against age-related cognitive decline, Alzheimer
      's disease, and Parkinson's disease.
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ANSWER 17 OF 20
                    MEDLINE on STN
                                                     DUPLICATE 4
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ACCESSION NUMBER: 2003386318 MEDLINE PubMed ID: 12923774 DOCUMENT NUMBER:

A panel of cerebrospinal fluid potential biomarkers for the TITLE:

diagnosis of Alzheimer's disease.

AUTHOR: Carrette Odile; Demalte Isabelle; Scherl Alexander;

Yalkinoglu Oezkarn; Corthals Garry; Burkhard Pierre;

Hochstrasser Denis F; Sanchez Jean-Charles

Biomedical Proteomics Research Group, Central Clinical CORPORATE SOURCE:

Chemistry Laboratory, Geneva University Hospital, 24 rue

Micheli-du-Crest, CH-1211 Geneva 14, Switzerland. Proteomics, (2003 Aug) Vol. 3, No. 8, pp. 1486-94.

Journal code: 101092707. ISSN: 1615-9853.

Germany: Germany, Federal Republic of PUB. COUNTRY: Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

LANGUAGE: English

Priority Journals FILE SEGMENT:

ENTRY MONTH: 200405

SOURCE:

ENTRY DATE: Entered STN: 19 Aug 2003

> Last Updated on STN: 18 May 2004 Entered Medline: 17 May 2004

The diagnosis of Alzheimer's disease (AD), the most common form AB

of dementia in the general population, usually relies upon the presence of typical clinical features and structural changes on brain magnetic resonance imaging. Over the last decade, a number of biological abnormalities have been reported in the cerebrospinal fluid (CSF) of AD patients, in particular altered levels of the tau protein and the 1-42 fragment of the amyloid precursor protein. These, however, have not yet proved sensitive and specific enough to be included in the diagnostic criteria for AD, leaving plenty of room for the search of novel biomarkers. The present study describes the analysis of CSF polypeptides by a protein-chip array technology called surface enhanced laser desorption/ionization-time of flight-mass spectrometry (SELDI-TOF-MS). Using this approach, we detected statistically significant quantitative differences (p < 0.05) regarding four overexpressed and one underexpressed polypeptides in the CSF of AD patients as compared to healthy controls. Four of them were further purified by strong anionic exchange chromatography (SAX) and identified by MS analysis as cystatin C, two beta-2-microglobulin isoforms, an unknown 7.7 kDa polypeptide, and a 4.8 kDa VGF polypeptide. The combination of the five polypeptides for the diagnosis of AD allowed to classified six AD patients out of the nine included in this study and all the ten controls, which means in this small cohort that the specificity and sensitivity are 100% and 66%, respectively. This study, based on the protein-chip array technology, demonstrates the presence in the CSF of novel potential biomarkers for AD, which may be used for the diagnosis and perhaps the assessment of the severity and progression of the disease.

L3 ANSWER 18 OF 20 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: . 2002:793918 CAPLUS

DOCUMENT NUMBER: 137:306353

TITLE: Neuroendocrine-specific protein VGF-derived

peptides VGFARP and their use in treatment

and diagnosis of dementia

INVENTOR(S): Lamping, Norbert; Zucht, Hans-Dieter; Heine, Gabriele;

Juergens, Michael; Hess, Ruediger; Selle, Hartmut;

Kellmann, Markus

PATENT ASSIGNEE(S): Biovision AG, Germany

SOURCE: PCT Int. Appl., 75 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: German

FAMILY ACC. NUM. COUNT: 1

PATENT NO.						DATE		APPLICATION NO.					DATE				
WO WO	D 2002082075     A2     200       D 2002082075     A9     200						2002	0021017 WO 2002-DE1376 2002040 0021219								108	
		AE, CO, GM, LS,	AG, CR, HR, LT,	AL, CU, HU, LU,	AM, CZ, ID, LV,	AT, DE, IL, MA,	AU, DK, IN, MD, SE,	AZ, DM, IS, MG,	DZ, JP, MK,	EC, KE, MN,	EE, KG, MW,	ES, KP, MX,	FI, KR, MZ,	GB, KZ, NO,	GD, LC, NZ,	GE, LK, OM,	GH, LR, PH,
	RW:	UA, GH, KG, GR,	UG, GM, KZ, IE,	US, KE, MD, IT,	UZ, LS, RU, LU,	VN, MW, TJ, MC,	YU, MZ, TM, NL, NE,	ZA, SD, AT, PT,	ZM, SL, BE, SE,	ZW SZ, CH, TR,	TZ,	UG, DE,	ZM, DK,	ZW, ES,	AM, FI,	AZ, FR,	BY, GB,
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EP		AT,	BE,	CH,	DE,	DK,	20040 ES, RO,	FR,	GB,	GR,	IT,						
	20045 20045	53125	50		T	•	2004	1014	Ċ	JP 20	002-5					0204 0310	

AB The invention provides Alzheimer's disease-associated peptides and methods for their detection and use in diagnosis and treatment of Alzheimer's disease. The peptides are proteolytic cleavage products of neuroendocrine-specific protein VGF. Changes in the concns. of said peptides indicate Alzheimer's disease, and the direction of the change in concentration is specific for each peptide. Alzheimer's disease is detected by identifying the peptides individually or in groups. The invention can also be used to control the course of Alzheimer's disease, for the prognosis thereof and for the development of therapeutic agents to combat the same.

L3 ANSWER 19 OF 20 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2002:937303 CAPLUS

DOCUMENT NUMBER: 138:20443

TITLE: Endocrine disruptor screening using DNA chips of

endocrine disruptor-responsive genes

INVENTOR(S): Kondo, Akihiro; Takeda, Takeshi; Mizutani, Shigetoshi;

Tsujimoto, Yoshimasa; Takashima, Ryokichi; Enoki,

Yuki; Kato, Ikunoshin

PATENT ASSIGNEE(S): Takara Bio Inc., Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 386 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
				-	
JP 2002355079	A	20021210	JP 2002-69354		20020313
PRIORITY APPLN. INFO.:			JP 2001-73183	Α	20010314
			JP 2001-74993	Α	20010315
			JP 2001-102519	Α	20010330

AB A method and kit for detecting endocrine-disrupting chems. using DNA microarrays are claimed. The method comprises preparing a nucleic acid sample containing mRNAs or cDNAs originating in cells, tissues, or organisms which have been brought into contact with a sample containing the endocrine disruptor. The nucleic acid sample is hybridized with DNA microarrays having genes affected by the endocrine disruptor or DNA fragments originating in these genes have been fixed. The results obtained are then compared with the results obtained with the control sample to select the gene affected by the endocrine disruptor. Genes whose expression is altered by tri-Bu tin, 4-octaphenol, 4-nonylphenol, di-N-Bu phthalate, dichlorohexyl phthalate, octachlorostyrene, benzophenone, diethylhexyl phthalate, diethylstilbestrol (DES), and 17- $\beta$  estradiol (E2), were found in mice by DNA chip anal.

L3 ANSWER 20 OF 20 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2001:300737 CAPLUS

DOCUMENT NUMBER: 134:321579

TITLE: Modulation of cell phenotype by transformation with

cAMP responsive element-binding proteins

INVENTOR(S): Reusch, Jane E.; Klemm, Dwight J.

PATENT ASSIGNEE(S): University Technology Corporation, USA; National

Jewish Medical and Research Center; U.S. Government as

Represented by the Department of Veterans Affairs

SOURCE: PCT Int. Appl., 155 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT NO.				KIND DATE		APPLICATION NO.					DATE						
	2001				A2		2001		1	WO 2	000-1	US28:	316		2	0001	012
WO	2001	0290	62		A3		2001	0913									
WO	2001	0290	62		A9		2002	8080	•								
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		CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EE,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,
		HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	ΚP,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,
		LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	PL,	PT,	RO,	RU,
		SD,	SE,	SG,	SI,	SK,	SL,	ΤJ,	TM,	TR,	TT,	TZ,	UA,	UG,	UZ,	VN,	YU,
		ZA,	ZW,	AM,	ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	TJ,	TM					
	RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	ŪĠ,	ZW,	ΑT,	BE,	CH,	CY,
		DE,	DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,
		CF,	CG,	CI,	CM,	GA,	GN,	GW,	ML,	MR,	NE,	SN,	TD,	TG			
AU	2001	0108	29		A		2001	0430	;	AU 2	001-	1082	9		2	0001	012
US	2004	0974	54		A1		2004	0520	1	US 2	003-	4315	98		2	0030	506
PRIORIT	Y APP	LN.	INFO	. :					1	US 1	999-	4200	60	Ž	A 1	9991	018
									1	WO 2	000-1	US28:	316	7	W 2	0001	012

Described is a method for modulating the phenotype of a cell, and AΒ particularly, of a target cell in a patient who has or is at risk of developing a disease or condition in which is associated with dysregulation of cellular phenotype. The method includes administration of a recombinant nucleic acid mol. encoding a protein having cAMP responsive element-binding (CREB) biol. activity or dominant neg. CREB biol. activity to a patient, in such a manner that the protein is expressed in a target cell of a patient and is sufficient to modulate the phenotype of the target cell. CREB is necessary and sufficient to initiate adipocyte differentiation, based on its constitutive expression in 3T3-L1 fibroblasts prior to the induction of adipogenesis and throughout the differentiation process. Furthermore, both CREB phosphorylation and transcriptional activity are rapidly induced in 3T3-L1 fibroblasts by conventional differentiation-inducing agents, and CREB binds to and stimulates transcription from the promoters of several adipocyte-specific genes. Augmentation of CREB protein expression by adenovrial gene transfer at the time of angioplasty will promoter smooth muscle cell differentiation and thereby decrease post-angioplasty restenosis. Such a method is particularly useful in patients who have, or at risk of developing, diabetes, obesity, macrovascular disease, heart failure, osteoarthritis, and neural diseases and conditions.

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FILE CONTAINS CURRENT INFORMATION.
LAST RELOADED: Feb 16, 2007 (20070216/UP).

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ANSWER 1 OF 3 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2002:793918 CAPLUS

DOCUMENT NUMBER:

137:306353

TITLE:

Neuroendocrine-specific

protein VGF-derived peptides

VGFARP and their use in treatment and

diagnosis of dementia

INVENTOR(S):

Lamping, Norbert; Zucht, Hans-Dieter; Heine, Gabriele;

Juergens, Michael; Hess, Ruediger; Selle, Hartmut;

Kellmann, Markus

PATENT ASSIGNEE(S):

Biovision AG, Germany

SOURCE:

PCT Int. Appl., 75 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

German

FAMILY ACC. NUM. COUNT: 1

PATENT NO.	KIND DAT	E APP	LICATION NO.	DATE
WO 2002082075	A2 200	21017 WO	2002-DE1376	20020408
WO 2002082075	A9 200	21219		
WO 2002082075	A3 200	30821		
W: AE, AG, AL,	AM, AT, AU	J, AZ, BA, BB	, BG, BR, BY, B	Z, CA, CH, CN,
CO, CR, CU,	CZ, DE, DK	C, DM, DZ, EC	, EE, ES, FI, G	B, GD, GE, GH,
GM, HR, HU,	ID, IL, IN	I, IS, JP, KE	, KG, KP, KR, K	Z, LC, LK, LR,

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           RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
      CA 2446886
                                  A1
                                           20021017
                                                           CA 2002-2446886
                                                                                          20020408
                                           20040102
                                                          EP 2002-742678
                                                                                          20020408
      EP 1373905
                                  A2
                AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
                 IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
      JP 2004531250
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                                           20041014
                                                           JP 2002-579794
                                                                                          20020408
                                           20040722
                                                           US 2003-680087
                                                                                          20031006
      US 2004142388
                                  A1
PRIORITY APPLN. INFO.:
                                                           DE 2001-10117431
                                                                                      A 20010406
                                                           WO 2002-DE1376
                                                                                      W 20020408
      The invention provides Alzheimer's disease-associated peptides and methods
```

AB The invention provides Alzheimer's disease-associated peptides and methods for their detection and use in diagnosis and treatment of Alzheimer's disease. The peptides are proteolytic cleavage products of neuroendocrine-specific protein VGF.

Changes in the concns. of said peptides indicate Alzheimer's disease, and the direction of the change in concentration is specific for each peptide. Alzheimer's disease is detected by identifying the peptides individually or in groups. The invention can also be used to control the course of Alzheimer's disease, for the prognosis thereof and for the development of therapeutic agents to combat the same.

L2 ANSWER 2 OF 3 MEDLINE on STN DUPLICATE 1

ACCESSION NUMBER: 2001473899 MEDLINE DOCUMENT NUMBER: PubMed ID: 11339279

TITLE: Peptide repertoire of human cerebrospinal fluid: novel

proteolytic fragments of neuroendocrine proteins.

AUTHOR: Stark M; Danielsson O; Griffiths W J; Jornvall H; Johansson

J

CORPORATE SOURCE: Department of Medical Biochemistry and Biophysics,

Karolinska Institutet, Stockholm, Sweden.

SOURCE: . Journal of chromatography. B, Biomedical sciences and

applications, (2001 Apr 25) Vol. 754, No. 2, pp. 357-67.

Journal code: 9714109. ISSN: 1387-2273.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200108

ENTRY DATE: Entered STN: 27 Aug 2001

Last Updated on STN: 27 Aug 2001` Entered Medline: 23 Aug 2001

AΒ Polypeptides in human cerebrospinal fluid (CSF), isolated by phase separation in chloroform-methanol-water and reversed-phase HPLC, were characterised by sequence analysis and mass spectrometry. This identified the presence of peptide fragments of testican, neuroendocrine specific protein VGF, neuroendocrine protein 7B2, chromogranin B/secretogranin I, chromogranin A, osteopontin, IGF-II E-peptide and proenkephalin. The majority of these fragments were generated by proteolysis at dibasic sites, suggesting that they are derived by activities related to prohormone convertase(s). Several of the fragments have previously not been detected, and their functions in CSF or elsewhere are unknown. A characteristic feature of all these fragments is a very high content of acidic residues, in particular glutamic acid. In addition to the fragments of neuroendocrine proteins, endothelin-binding receptor-like protein 2, ribonuclease 1, IGF-binding protein 6, albumin, alphal-acid glycoprotein 1, prostaglandin-H2 D-isomerase, apolipoprotein Al, transthyretin, beta2-microglobulin, ubiquitin, fibrinopeptide A, and C4A anaphylatoxin were found.

L2 ANSWER 3 OF 3 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1997:702966 CAPLUS

DOCUMENT NUMBER: 128:44439

TITLE: Cloning, structural organization analysis, and

chromosomal assignment of the human gene for the

-1.56

-1.56

neurosecretory protein VGF

AUTHOR(S): Canu, Nadia; Possenti, Roberta; Ricco, Angela Serena;

Rocchi, Mariano; Levi, Andrea

CORPORATE SOURCE: Dip. Med. Sperimentale Sci. Biomed., Seconda Univ.

Roma Tor Vergata, Rome, 00173, Italy

SOURCE: Genomics (1997), 45(2), 443-446

CODEN: GNMCEP; ISSN: 0888-7543

PUBLISHER: Academic
DOCUMENT TYPE: Journal
LANGUAGE: English

The Vgf gene was originally identified as a 2.7-kb cDNA fragment isolated from nerve growth factor-treated PC12 cells by differential display against PC12 cells. It is transcribed solely in subpopulations of neuroendocrine cells in vivo and it is induced by neurotrophins in target cells in vitro. The single-copy human VGF gene was isolated from a genomic library. The gene spans approx. 6 kb and contains two exons. The entire VGF protein is encoded by exon 2, while exon 1 contains only 5'-untranslated sequence. The structural organization of the human gene is similar to that described for the rat Vgf gene and both the translated and the untranslated regions show a high degree of sequence homol. to the rat gene. Northern blot anal. revealed a single transcript of approx. 2.7 kb that was detected only in mRNA prepns. from brain. The gene was assigned to chromosome 7g22 by fluorescence in situ hybridization.

REFERENCE COUNT: 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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